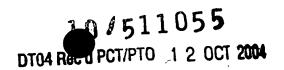


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## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

IN RE	E: Application of Steven Gerard BLANCHARD et al.					
Serial	No.: To be assigned	Art Unit:				
Filing	Date: Concurrently herewith	Examiner:				
For:	Compositions and Methods for Evaluating and Designing Nuclear Receptor Ligands that Modulate Co-regulator Affinity					
P.O. I	nissioner for Patents Box 1450 ndria, VA 22313-1450					
	INFORMATION DISCLO	OSURE STATEMENT				
	cants request that the references identified on Form PT iner and officially made of record in accordance with t					
[	[X] Copies of the references listed on the attached form:  [] Copies of the references were submitted in parent at a copy of the International Search Report which is:  [] PCT/US03/11055 is submitted had International Search Report are listed on the attached understand that copies have been supplied to the U.	pplication Serial No (37 CFR 1.98(d)) sued on International Application No. erewith. All of the publications cited in the ed form PTO-1449 as item no. 2 and Applicants				
A. [2	filing date of the above application or date	emitted herewith is being filed within three months of the cof entry into the national stage of an international first Office action on the merits, whichever event				
Ĩ	] The Information Disclosure Statement	submitted herewith is being filed before the mailing a Request For Continued Examination under 37				
В. [	filing date of the above application or the date	tted herewith is being filed <b>after</b> three months of the of entry into the national stage as set forth in § 1.491 of g date of the first Office Action on the merits, whichever e of either:				
į	Applicant hereby certifies that each item of inf Statement was cited in a communication from a application not more than three months prior to					

Applicant elects the option to pay the fee set forth in 37 CFR 1.17(p) for submission of an Information Disclosure Statement under § 1.97(c) (\$180.00).

[]

C. [] The Information Disclosure Statement transmitted herewith is being filed after a final action under § 1.113, or a notice of allowance under § 1.311, whichever occurs first, but before the payment of the issue fee. Also enclosed is a copy of the International Search Report which Issued on International Publication No.

In accordance with the requirements of 37 CFR 1.97(d):

- Applicant hereby certifies that each item of information contained in this Information Disclosure Statement was cited in a communication from a foreign patent office in a counterpart foreign application not more than three months prior to the filing of this statement. [or]
- Applicant hereby certifies that no item of information contained in this Information Disclosure Statement was cited in a communication from a foreign patent office in a counterpart foreign application, and, to my knowledge after making reasonable inquiry, no item of information contained in this Information Disclosure Statement was known to any individual designated in § 1.56(c) more than three months prior to the filing of this statement; and
- [ ] The petition fee set forth in § 1.17(i)(1) (\$180.00) is submitted herewith.
- [X] Please charge any required fees to Deposit Account No.07-1392.

[ ] A duplicate copy of this paper is attached.

Respectfully Submitted,

Michael M. Conger Attorney of Record: Registration No. 43,562

Date: October 1, 2004

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		Bourguet et al., "Nuclear receptor ligand-binding domains: three-dimensional structures, molecular interactions and pharmacological implications," <i>Trends in Pharmacological Sciences</i> <b>21(10)</b> :381-388 (Oct. 2000).								
	4.	Bramlett et al., "Ligands specify coactivator nuclear receptor (NR) box affinity for estrogen receptor subtypes," <i>Molecular Endocrinology</i> <b>15(6)</b> :909-922 (Jun. 2001).								
	5.	Brown et al., "A reverses the diabe								
	6.	Brzozowski et al. 389(6652):753-75	, "Molecular bas							
	7.	Caira et al., "Clor Biological Chemi	ning and characte	erization of	RAP250, a nove	el nuclear recepto	r coactivator,"	Journal of		
	8.	Chen et al., "A tra				nuclear hormone	receptors," λ	lature		
		377(6548):454-45								
	9.	Cobb et al., "N-(2 optimization of the	ne N-aryl substitu	uent," Journ	al of Medicinal	Chemistry 41(25	):5055-5069 (	Dec. 1998)		
	10.	Cohen et al., "The nuclear corepressors recognize distinct nuclear receptor complexes," <i>Molecular Endocrinology</i> <b>14(6)</b> :900-914 (Jun. 2000).								
	11.									
	12.									
	13.									
	the interactions of retinoid X receptor heterodimers with ligands, coactivators, and corepressors," <i>Molecula &amp; Cellular Biology</i> 17(4):2166-2176 (Apr. 1997).									
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	14.	Dowell et al., "p300 functions as a coactivator for the peroxisome proliferators-activated receptor alpha," Journal of Biological Chemistry 272(52):33435-33443 (Dec. 1997).									
	15.	Gampe et al., "Asymmetry in the PPARgamma/RXRalpha crystal structure reveals the molecular basis of heterodimerization among nuclear receptors," <i>Molecular Cell</i> 5(3):545-555 (Mar. 2000).									
<u> </u>	16.	Gee et al., "Coac						gens and			
	<u> </u>	antiestrogens wit	h the estrogen rec	ceptor," Mol	lecular Endocrii	nology <b>13(11)</b> :19	12-1923 (Nov	. 1999).			
	17.	Grignani et al., "I promyelocytic le					one deacetylas	e in			
	18.	Henke et al., "N-	(2-Benzoylpheny	l)-L-tyrosin	e PPARgamma	agonists. 1. Disc					
		Henke et al., "N-(2-Benzoylphenyl)-L-tyrosine PPARgamma agonists. 1. Discovery of a novel series of potent antihyperglycemic and antihyperlipidemic agents, Journal of Medicinal Chemistry 41(25):5020-5036 (Dec. 1998).									
	19.		Horlein et al., "Ligand-independent repression by the thyroid hormone receptor mediated by anuclear								
	20.	receptor co-repressor," <i>Nature</i> <b>377(6548)</b> :397-404 (Oct. 1995).  Hu et al., "The CoRNR motif controls the recruitment of corepressors by nuclear hormone receptors," <i>Nature</i>									
		402(6757):93-96		5.5		F. 3555.5 OJ Mach		eropioio, i	········		
	21.			nember of th	e steroid hormo	ne receptor supe	rfamily by per	oxisome			
		Issemann et al., "Activation of a member of the steroid hormone receptor superfamily by peroxisome proliferators," <i>Nature</i> <b>347(6294)</b> :645-650 (Oct. 1990).									
	22.	Kliewer et al., "Differential expression and activation of a family of murine peroxisome proliferators-activated receptors," <i>Proc. Nat'l. Acad. Sci. USA</i> 91(15):7355-7359 (Jul. 1994).									
	23.	Lavinsky et al., "Diverse signaling pathways modulate nuclear receptor recruitment of N-CoR and SMRT									
	24.	complexes Proc. Nat'l. Acad. Sci. USA 95(6):2920-2925 (Mar. 1998).									
	27.	Lee et al., "A nuclear factor, ASC-2, as a cancer-amplified transcriptional coactivator essential for ligand-dependent transactivation by nuclear receptors in vivo," Journal of Biological Chemistry 274(48):34283-									
	34293 (Nov. 1999).								_		
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~	DTHER DOCUMENTS (Including Author, Title, Journal-Date, Page Number, Etc.) (Cont'd)  25. Lehmann et al., "An antidiabetic thiazolidinedione is a high affinity ligand for peroxisome proliferators-activated receptor gamma (PPAR gamma)," Journal of Biological Chemistry 270(22):12953-12956 (Jun. 1995).									
	26.	Nagy et al., "Mecl Development 13(2			ng and release f	rom nuclear horm	one receptors	," Genes &		
	27.	Nolte et al., "Liga gamma," Nature 3	nd binding and co	o-activator 3 (Sep. 19	98).			•		
	28.	Norris et al., "Pep								
	29.	Oliver et al., "A se cholesterol transpe						reverse		
-	30.	Perissi et al., "Mo 13(24):3198-3208	lecular determina					Development		
	31.	Pike et al., "Struct	ture of the ligand-				in the presence	of a partial		
	32.	agonist and a full antagonist," <i>EMBO Journal</i> 18(17):4608-4618 (Sep. 1999).:  Puigserver et al., "A cold-inducible coactivator of nuclear receptors linked to adaptive thermogenesis," <i>Cell</i> 92(6):829-839 (Mar. 1998).								
	33.	Robyr et al., "Nuclear hormone receptor coregulators in action: diversity for shared tasks," Molecular								
	34.	,								
interaction by tamoxifen," Cell 95(7):927-937 (Dec. 1998.)										
	35. Warnmark et al., "Differential recruitment of the mammalian mediator subunit TRAP220 by estrogen receptors ERalpha and ERbeta," J. Biol. Chem. 276(26):23397-23404 (Jun. 2001).									
	Webb et al., "The nuclear receptor corepressor (N-CoR) contains three isoleucine motifs (I/LXXII) that ser as receptor interaction domains (IDs)," <i>Molecular Endocrinology</i> 14(12):1976-1985 (Dec. 2000).									
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	37.	THER DOCUMENTS (Including Author, Title, Journal-Date, Page Number, Etc.) (Cont'd)  Willson et al., "The PPARs: from orphan receptors to drug discovery," Journal of Medicinal Chemistry  43(4):527-550 (Feb. 2000).								
	38.	Wong et al., "Structure-function evaluation of ER alpha and beta interplay with SRC family coactivators. ER selective ligands," <i>Biochemistry</i> <b>40(23)</b> :6756-6765 (Jun. 2001).								
	39.									
	40.	activated receptor	s, " Proc. Nat'l	Acad. Sci. U	inding selectivity l USA <b>98(24)</b> :13919-	-13924 (Nov. 2	2001).			
	41. Xu et al., "Structural basis for antagonist-mediated recruitment of nuclear co-repressors by PPARα," National 415(6873):813-817 (Feb. 2002).									
	42. Yamamoto et al., "The Tamoxifen-responsive estrogen receptor alpha mutant D351Y shows reduced Tamoxifen-dependent interaction with corepressor complexes," <i>Journal of Biological Chemistry</i> 276(46):42684-42691 (Nov. 2001).									
	43.	receptors and tran	scriptional corep	ressors," M		ology 11(4):47	0-480 (Apr. 1	997).		
	44.	receptors and transcriptional corepressors," <i>Molecular Endocrinology</i> 11(4):470-480 (Apr. 1997).  44. Yuan et al., "The TRAP220 component of a thyroid hormone receptor-associated protein (TRAP) coactivator complex interacts directly with nuclear receptors in a ligand-dependent fashion," <i>Proc. Nat'l Acad. Of Sci. USA</i> 95(14):7939-7944 (Jul. 1998).								
	45.	Zhang et al., "A n	uclear receptor c	orepressor	modulates transcri			-occupied		
	46.	steroid hormone receptor," <i>Molecular Endocrinology</i> 12(4):513-524 (Apr. 1998).  46. Zhu et al., "Cloning and identification of mouse steroid receptor coactivator-1 (mSRC-1), as a coactivator of peroxisome proliferators-activated receptor gamma," <i>Gene Expression</i> 6(3):185-195 (1996).								
	<ul> <li>47. Zhu et al., "Isolation and characterization of PBP, a protein that interacts with peroxisome proliferators-activated receptor," <i>Journal of Biological Chemistry</i> 272(41):25500-25506 (Oct. 1997).</li> </ul>									
	48. Zhu et al., "Isolation and characterization of peroxisome proliferators-activated receptor (PPAR interacting protein (PRIP) as a coactivator for PPAR," Journal of Biological Chemistry 275(18):13510-13516 (May									
	2000).									
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